RESEARCH PAPER

Maternal Age at Childbirth and Perinatal and Under-five Mortality in a Prospective Birth Cohort from Delhi

^{*#}Sikha Sinha, ^{\$}Abha Rani Aggarwal, [‡]Clive Osmond, [‡]Caroline HD Fall, [^]Santosh K Bhargava and *Harshpal Singh Sachdev

From ^{*}Sitaram Bhartia Institute of Science and Research, New Delhi, India; [#]University School of Medicine and Para-medical Health Sciences, Guru Gobind Singh Indraprastha University, Delhi, India; ^{\$}National Institute of Medical Statistics, Indian Council of Medical Research, New Delhi, India; [‡]MRC Lifecourse Epidemiology Unit, University of Southampton, Southampton, UK; and [^]Sunder Lal Jain Hospital, New Delhi, India.

Correspondence to: Prof HPS Sachdev, Senior Consultant Pediatrics and Clinical Epidemiology, Sitaram Bhartia Institute of Science and Research, B-16 Qutab Institutional Area, New Delhi 110016, India. hpssachdev@gmail.com Received: April 28, 2016; Initial review: June 01, 2016; Accepted: July 07, 2016.

Objective: To evaluate the relationship between maternal age at child birth, and perinatal and under-five mortality.

Design: Prospective birth cohort.

Setting: Urban community.

Participants: 9169 pregnancies in the New Delhi Birth Cohort resulted in 8181 live births. These children were followed for survival status and anthropometric measurements at birth (+3 days), 3,6,9 and 12 months (7 days), and every 6 months thereafter until 21 years age. Information on maternal age at child birth and socio-demographic profile was also obtained.

Outcome measures: Offspring mortality from 28 weeks gestation till 5 years age.

Results: Offspring mortality (stillbirths -5 years; n=328) had a U-shaped association with maternal age (P<0.001). Compared to the reference group (20-24 years), younger (≤ 19 years) and older

(≥ 35 years) maternal ages were associated with a higher risk of offspring mortality (HR: 1.68; 95% CI 1.16, 2.43 and HR 1.48; 95% CI 1.01, 2.16, respectively). In young mothers, the increased risk persisted after adjustment for socio-economic confounders (maternal education, household income and wealth; HR 1.51; 95% CI 1.03, 2.20) and further for additional behavioral (place of delivery) and biological mediators (gestation and birthweight) (HR 2.14; 95% CI 1.25, 3.64). Similar associations were documented for post-perinatal deaths but for perinatal mortality the higher risk was not statistically significant (P >0.05). In older mothers, the increased mortality risk was not statistically significant (P >0.05) after adjustment for socio-economic confounders.

Conclusion: Young motherhood is associated with an increased risk of post-perinatal mortality and measures to prevent early childbearing should be strengthened.

Key Words: Child mortality, Risk factors, Teenage pregnancy.

Published online: July 10, 2016. PII:S097475591600015

eduction of under-five child mortality, the target of Millennium Development Goal 4 (MDG 4), has shown remarkable progress globally since 1990, with the highest average annual reduction rate of 4% during 2005-2013 [1]. Sub-Saharan Africa and South Asia continue to have the highest under-five mortality burden; India had 49 under-five deaths per 1000 live births in 2013 [2], and is lagging behind the committed target [3,4]. Perinatal mortality, which includes stillbirth, has received much less global attention despite being most common in low- and middle-income countries (LMIC) [5], and has declined at a slower rate than under-five mortality.

Current interventions for improving child health and survival focus primarily on medical aspects including immunization, and improving access to healthcare and illness management, eventhough social factors are also important. Optimal maternal age at child bearing is one such undervalued factor [6]. Early marriage and childbearing are still quite prevalent in India, especially in rural areas; 18% and 47% are married before 15 years and 18 years, respectively [7]. If extremes of maternal age contribute substantially to stillbirths and child mortality, ensuring an optimal age at childbirth merits greater priority as an intervention for accelerating progress.

Accompanying Editorial: Pages 868-69.

Cross-sectional data suggest that children born to mothers <20 years of age are at increased risk for perinatal, neonatal and under-five child mortality [8-12]. However, this existing evidence has important methodological limitations. There is scant data from longitudinal cohorts in LMIC [13] exploring the association between maternal age at childbirth and mortality, particularly in relation to stillbirths. We, therefore, evaluated the relationship of maternal age with perinatal and under-five mortality in the New Delhi Birth Cohort (NDBC), using appropriate statistical techniques and adjustment for confounders and mediators.

METHODS

The NDBC was drawn from a population of 119,799 living in a 12 km² area of south Delhi during 1969-72 [14,15]. 20,755 married women of reproductive age were recruited and followed regularly every other month to record menstrual dates. During recruitment, a social worker obtained information on maternal schooling and age, household structure including family income, number of family-members, ownership and type of residence, and sanitation and water supply facilities. Women who became pregnant were followed every two months initially and on alternate days from the 37th week of gestation to determine the pregnancy outcome. There were 9169 pregnancies, resulting in 8181 live births. Survival status and anthropometric measurements (length and weight) of these babies were recorded within 72 hrs of birth, at the ages of 3, 6, 9 and 12 months (± 7) days) and every 6 months until 21 years by trained personnel.

Statistical analysis: From the available data, mortality could be categorized as perinatal (28 weeks gestation to 6 postnatal days), late neonatal (7-28 days), postneonatal to infant (29 days-1 year), and thereafter at yearly intervals until 5 years. However, due to small numbers in each of these categories, we used the following categories in our analysis: (*i*) all deaths between 28 weeks of gestation and five years of age (including stillbirths), (*ii*) perinatal mortality (28 weeks of gestation-6 days), and (*iii*) post-perinatal mortality (7 days-5 years age).

Data analysis was performed using SPSS version 20.0. Student's *t*-test and Chi square test were used to compare descriptive statistics between alive and dead cases. Associations of maternal age at birth with mortality were determined using Cox Proportional Hazard Model [16]. Maternal age was initially used in a continuous format and the quadratic term was used to assess the non-linear associations. Subsequently, it was divided into five groups (\leq 19, 20-24, 25-29, 30-34 and \geq 35 years) with 20-24 years (maximum sample size) as the reference category.

The associations between maternal age and offspring mortality were evaluated in a stepwise manner. Crude analyses adjusted for the child's sex, followed by adjustment for confounders, and later for additional mediators. We included only those potential confounders and mediators, which were significantly (P<0.05) different between children who survived and those who died. Confounders included for adjustment were socioeconomic factors (maternal education, per capita annual household income and household wealth). Household wealth scores were derived from the 1st principal component [17] for the combination of type of housing and ownership, sanitation, water supply and crowding (number of people/room); a higher score related to better wealth. The potential mediators available and considered for additional adjustment were behavioural (place of delivery and breastfeeding status), and biological (birth weight and gestation). As breastfeeding status was relevant only for the post-perinatal deaths, it was not included. The final primary analyses models were: (i) Model 1- adjusted for sex, (ii) Model 2adjusted for sex and socio-economic confounders (maternal education, household income and household wealth); and (iii) Model 3- adjusted for sex, socioeconomic confounders and mediators (place of delivery, gestation and birth weight). A sensitivity analysis was also performed on Model 3 with additional adjustments for breastfeeding status (only for post-perinatal deaths). Linear and quadratic associations between maternal age and socio-economic confounders and mediators were also analyzed.

RESULTS

At the time of recruitment in 1969-1972, 60 percent of cohort families had an income above 50 rupees per month (national average, 28) and only 15 percent of parents were illiterate (national average, 66). Nevertheless, 43% of families lived in one room. Hindus were the majority religious group (84%) [15]. Information on maternal age at child birth was available for 5886 subjects (mean (SD) age 25.9 (5.3) years). All of them were married and 67% of them living in masonry buildings with good water supply and sanitation facilities. Only 31.5% of the mothers had received 10 or more years of education.

There were 328 deaths reported up to 5 years of age including stillbirths, with no significant sex differences (*Table I*). Most deaths (84%) had occurred by 1 year of age, with neonatal to infant (41.1%), perinatal (29.0%) and late neonatal (13.7%) deaths being the major contributors. Demographic and birth characteristics among those censored (alive) and those who died are compared in *Web Table I*. Considering all deaths, children who had died were born smaller and at an earlier gestation than survivors. Their mothers had less education and poorer housing, water supply and sanitation facilities, and lower per capita annual household income and household wealth scores.

TABLE I SEX-WISE MORTALITY DISTRIBUTION

Mortality period	Male	Female	Total
Perinatal (28 wk gestation-6 d)	52 (33.3)	43 (25.0)	95 (29.0)
Late neonatal (7 d-28 d)	21 (13.5)	24 (13.9)	45 (13.7)
Post-neonatal infant (29 d-1 y)	64 (41.0)	71 (41.3)	135 (41.1)
1-2 у	10 (6.4)	23 (13.4)	33 (10.0)
2-3 у	5 (3.2)	5 (2.9)	10 (3.1)
3-5 у	4 (2.6)	6 (3.5)	10 (3.1)
Total	156	172	328

All values in No.(%). No statistically significant sex differences.

However, there were no differences in mean maternal age at childbirth and birth order. An analysis restricted to post-perinatal and perinatal deaths, yielded similar findings. Predominant breastfeeding was nearly universal (98.9% at birth and 91.5% at 3 months) but practised more often in survivors. However, for perinatal deaths, the place of delivery and most of the socio-economic variables were not significantly different, except for household income and house-ownership.

All the socio-economic confounders (maternal education, household income and household wealth), and mediators (place of delivery, gestation and birthweight) had inverted U-shaped relationship with maternal age ($P \le 0.001$ for quadratic term) (*Web Table* II). Both younger and older age of mothers was associated with lower education, household income, wealth, birthweight and gestation, and less likely to deliver in the healthcare services. Maternal age was unrelated to breastfeeding status.

Offspring mortality (stillbirths – 5 years) had a significant U-shaped relationship with maternal age (P<0.001), which persisted after adjustment for socioeconomic status confounders (P=0.003) and mediators (P=0.018) (**Web Table III**). There were similar associations, of borderline significance in the mediatoradjusted model (P=0.07), for post-perinatal deaths. However, for perinatal deaths there was no evidence of a significant (P>0.05) quadratic association.

All deaths (stillbirths and mortality till five years of age): **Table II** depicts the risk of offspring mortality across the five maternal age groups. In comparison to mothers aged 20-24 years, younger (\leq 19 years) and older (\geq 35 years) maternal ages were associated with higher offspring mortality (stillbirth – 5 years) (HR:1.68; 95% CI 1.16, 2.43 and HR 1.48; 95% CI 1.01, 2.16, respectively). After adjustment for socio-economic confounders, this higher risk persisted for younger mothers (HR 1.51; 95% CI 1.03, 2.20) but not for older

mothers (HR 0.99; 95% CI 0.66, 1.48). On further adjustment for mediators, offspring of both younger and older mothers had a higher risk of mortality (HR 2.14; 95% CI 1.25, 3.64 and HR 1.74; 95% CI 1.02, 2.97, respectively). In order to estimate the change in effect size of the association with additional confounder and mediator adjustments (which led to reductions in sample size), models 1 and 2 were run on the available sample for the fully adjusted model 3 (*Fig.* 1). The hazard ratios for both younger and older mothers were sequentially attenuated from the crude to the fully adjusted models. The mothers available for fully adjusted model 3 (after reduction in sample size) were comparatively educated and had marginally higher household income and wealth score.

Post-perinatal or perinatal deaths: A similar pattern was found for post-perinatal mortality; the increased risk being statistically significant (P<0.05) for all three models in younger but not older mothers. The attenuation pattern was similar for perinatal deaths but the increased risk was not statistically significant. There were no instances for which the point estimate in one time interval was outside the 95% confidence for the other time interval, thereby suggesting that the effect sizes were similar or hazard was proportional in both perinatal and post-perinatal categories.

On sensitivity analyses (data not presented), the mortality risk for younger and older mothers remained similar after additional adjustments for birth-order (all three death categories) and breastfeeding (post-perinatal deaths).

DISCUSSION

In this prospective cohort study, offspring of young (<20 years) mothers had an increased risk of mortality from the perinatal period up to age five years, primarily after the early neonatal period. An apparently similar disadvantage in older (>35 years) mothers was principally a reflection of their adverse socio-economic profile.

Persistence of a higher overall mortality risk in children of young mothers, despite adjustments for confounders and mediators, suggests a causal relationship. Similar effects were evident for postperinatal deaths but not for perinatal mortality. This could either reflect a true biological difference or insufficient statistical power for the perinatal mortality component, which showed broadly similar associations (29-95 deaths in various models). The confounderadjusted association for post-perinatal mortality was further attenuated after the introduction of mediators and, except breastfeeding, the other three biological and

INDIAN PEDIATRICS

Variables	Model 1 Hazard ratio (95% CI) (P value)	Model 2 Hazard ratio (95% CI) (P value)	Model 3 Hazard ratio (95% CI) (P value)
All deaths			
Number of deaths/total sample (deaths + censored)	328/5886	316/5478	156/4154
Maternal age (years)			
-19	1.68 (1.16; 2.43) (0.006)	1.51 (1.03; 2.20) (0.033)	2.14 (1.25; 3.64) (0.005)
20-24	Reference	Reference	Reference
25-29	1.00 (0.76; 1.31) (0.982)	0.94 (0.71; 1.24) (0.655)	1.34 (0.88; 2.05) (0.178)
30-34	1.00 (0.71; 1.40) (0.990)	0.77 (0.54; 1.09) (0.140)	1.02 (0.59; 1.74) (0.956)
35+	1.48 (1.01; 2.16) (0.043)	0.99 (0.66; 1.48) (0.968)	1.74 (1.02; 2.97) (0.042)
Perinatal deaths			
Number of deaths/total sample (deaths + censored)	95/5886	91/5478	29/4154
Maternal age (years)			
-19	1.51 (0.78; 2.92) (0.219)	1.42 (0.72; 2.83) (0.312)	1.22 (0.32; 4.63) (0.775)
20-24	Reference	Reference	Reference
25-29	0.90 (0.54; 1.48) (0.667)	0.84 (0.50; 1.40) (0.498)	1.07 (0.40; 2.83) (0.891)
30-34	0.91 (0.49; 1.69) (0.759)	0.77 (0.40; 1.45) (0.410)	0.85 (0.22; 3.31) (0.817)
35+	0.93 (0.41; 2.09) (0.852)	0.61 (0.25; 1.49) (0.280)	1.73 (0.51; 5.90) (0.380)
Post-perinatal deaths			
Number of deaths/total sample (deaths + censored)	233/5483	225/5080	127/3894
Maternal age (years)			
-19	1.77 (1.13; 2.75) (0.012)	1.57 (1.00; 2.46) (0.052)	2.39 (1.33; 4.28) (0.003)
20-24	Reference	Reference	Reference
25-29	1.05 (0.75; 1.46) (0.790)	0.98 (0.70; 1.38) (0.911)	1.38 (0.86; 2.22) (0.180)
30-34	1.04 (0.70; 1.56) (0.845)	0.76 (0.50; 1.16) (0.209)	1.05 (0.59; 1.89) (0.862)
35+	1.73 (1.13; 2.67) (0.013)	1.14 (0.72; 1.79) (0.580)	1.69 (0.93; 3.08) (0.087)

TABLE II ASSOCIATION BETWEEN DIFFERENT MATERNAL AGE-GROUPS A	ND OFFSPRING MORTALITY
--	------------------------

Model 1: adjusted for sex; Model 2: adjusted for sex, socio-economic confounders (maternal education, household income and wealth); and Model 3: adjusted for sex, socio-economic confounders and biological mediators (place of delivery, gestation and birth weight).

behavioural factors (place of delivery, gestation and birth weight) were significantly related to young maternal age. The increased risk appears to be partly operating through lower birth weight and gestation [6], and less utilization of health care services (home delivery). These factors; however, are of limited relevance for the stillbirth component of perinatal mortality as the event is likely to determine the birth weight, gestation and access to health care rather than the converse. In contrast, the increased overall mortality risk in older mothers was not evident after socio-economic adjustments. Older maternal age may thus not biologically predispose the offspring to higher mortality, and older mothers are also likely to be more experienced in child care practices. In a recent meta-analysis of five

birth cohorts from LMIC (of which NDBC was one) children of older mothers had a higher risk of preterm birth, but had better nutritional status and schooling after similar confounder adjustment [6]. Older mothers available for the fully adjusted model 3 had higher education and wealth score, which along with a lower sample size could explain the observed statistically significant associations.

Earlier cross-sectional data, including pooled analyses from 118 demographic and health surveys conducted between 1990 and 2008 in 55 low and middle income countries (LMIC), also documented a higher risk of perinatal, neonatal, infant and under-five mortality in young mothers [8-12,18-23]. It is suggested that this risk

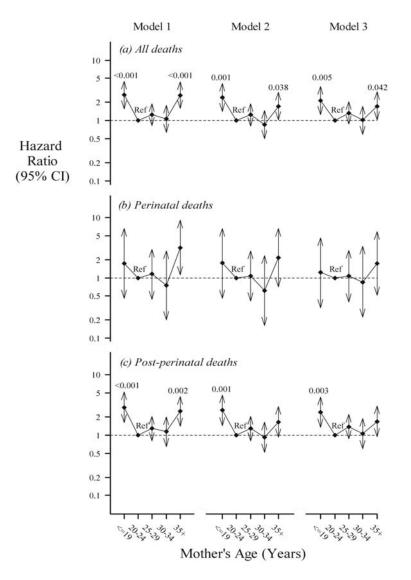


FIG. 1 Hazard ratio for mortality across different age groups of maternal age at childbirth (a) all deaths till five years including stillbirths (Number of deaths/total sample: 156/4154); (b) Perinatal deaths (Number of deaths/total sample: 29/4154); (c) Postperinatal deaths (Number of deaths/total sample: 127/3894). (Model 1: adjusted for sex; Model 2: further adjusted for socio-economic confounders and Model 3: further adjusted for mediators (type of delivery, gestation and birth weight). The bars represent 95% confidence interval for the hazard ratio and figures at the top of the bars are P value for significant age groups. Ref: Reference age group.

may operate through both biological and social mechanisms. Some studies also documented a higher risk in older mothers or J or U shaped association, particularly for unadjusted models [18,24]. However, this evidence has important limitations: (*a*) Crosssectional design and variation in context and time period; (*b*) Sub-optimal confounder adjustments; (*c*) Non-linear relationships have been rarely explored; and (*d*) Prospective data collection, to minimize bias, is mostly restricted to developed countries. Three population-based cohorts in Brazil (1982, 1993 and

2004) observed an increased risk of post-neonatal infant mortality (confounder adjusted OR 1.6; 95% CI 1.2, 2.1) in children of young (<20 years) mothers but not for stillbirths, perinatal deaths or neonatal mortality [13]. Further adjustment for mediating variables (place of delivery, gestation and birth weight) led to the disappearance of the excess of post-neonatal mortality. It was concluded that social and environmental factors may be more important than biological immaturity for this increased mortality. However, in our data, the increased risk for post-perinatal deaths persisted even after

INDIAN PEDIATRICS

WHAT IS ALREADY KNOWN?

Cross-sectional analyses, often with inadequate confounder adjustments, suggest that young motherhood is
associated with perinatal, neonatal and under-five mortality

WHAT THIS STUDY ADDS?

This prospective birth cohort data with confounder and mediator adjustments indicate that children of teenage
mothers are at an increased risk of post-perinatal mortality, and measures to prevent young motherhood should
be strengthened. An apparently similar disadvantage in older (>35 years) mothers is principally a reflection of
their adverse socio-economic profile.

confounder and mediator adjustment, suggesting a causal relationship. These observed differences, among other factors, could relate to contextual variability, baseline mortality risk, social characteristics of young mothers, social and health care support systems and methodological differences (surveillance versus prospective cohort follow up, including or excluding mothers \geq 30 years and restricting outcomes to infant or under-five mortality). We thus hypothesize that young maternal age predisposes the offspring to higher postperinatal mortality, which only partly operates through socio-economic deprivation and biological-behavioural mediators (lower birthweight and gestation, and poorer access to healthcare); the additional precise biological mechanisms need further exploration.

Strengths of our study are a large sample size, prospective community-based recording of confounders, mediators and outcomes until five years age from a South Asian setting, and appropriate analyses. The following limitations also merit consideration: (i) the relevance of four decades old data for contemporary programmes could be questioned. However, the findings have important programmatic implications for several regions in the country that even now have similar fertility, mortality, poor socio-economic, water supply and sanitation and health access indicators. Further, there was no evidence of secular changes in associations in data spread over 2-3 decades. [13,18]; (ii) data are missing for some variables; however, most of this pertains to mediators rather than confounders and this is a familiar scenario in large prospective cohort studies from LMIC; (iii) there may be some residual unadjusted confounding; (iv) a separate category of early neonatal deaths was not available for analysis. In community settings in India, it is challenging to discern a live newborn from a stillbirth within the first day of delivery.

Offspring of teenage mothers in LMIC not only have poorer child survival, but are also disadvantaged at birth and during childhood, and have reduced human capital [6]. Measures to prevent young motherhood are currently underrated as public health interventions; these should receive greater prominence and investments in the proposed child health and survival agenda [25]. Teenage marriages and pregnancies are declining in India [26,27]. However, as per latest national estimates, 32% of all women and 40% of those illiterate are married before 18 years [26]; the intervention thus still retains importance, particularly in rural and tribal regions. Further, greater care and support is necessitated for their vulnerable children in public health programs. It would be unethical to conduct randomized controlled trials on this subject. However, operational and behavioural research to prevent young motherhood in different contexts is desirable. Pooled analyses from recent similar cohorts in LMIC could confirm the utility of this intervention with improvements in access to health care.

In conclusion, children of teenage mothers are at an increased risk of post-perinatal mortality and measures to prevent young motherhood should be strengthened.

Contributors: SS, ARA, HPS, CHDF, SKB: conceptualised the study. SS, ARA, CO, HPS: analyzed the data. SS: drafted the initial manuscript. All authors contributed to the critical revision of the article.

Funding: Indian Council of Medical Research for supporting Ms. Sikha Sinha through the Senior Research Fellowship Scheme. The original cohort studies were supported by the National Center for Health Statistics, USA and the Indian Council of Medical Research.

Competing interest: None stated.

References

- 1. You D, Hug L, Ejdemyr S, Idele P, Hogan D, Mathers C, *et al.* Global, regional, and national levels and trends in under-5 mortality between 1990 and 2015,with scenario-based projections to 2030: A systematic analysis by the UN Inter-agency Group for Child Mortality Estimation. Lancet. 2015;386: 2275-86.
- 2. SRS Statistical Report, September 2014, Volume 49 No. 1. Available from: http://censusindia.gov.in/vital_statistics/ SRS_Bulletins/SRS%20Bulletin%20-Sepetember% 202014.pdf. Accessed March 30, 2016.
- 3. Paul VK, Sachdev HS, Mavalankar D, Ramachandran

P, Sankar MJ, Bhandari N, *et al.* Reproductive health, and child health and nutrition in India: meeting the challenge. Lancet. 2011;377:332-49.

- 4. IGME Report 2015 Child Mortality Final UNICEF. Available from: http://www.childmortality.org/files_v20/ download/IGME%20report%202015%20child% 20mortality%20final.pdf. Accessed March 30, 2016.
- Lawn JE, Blencowe H, Waiswa P, Amouzou A, Mathers C, Hogan D, *et al.*; Lancet Ending Preventable Stillbirths Series study group; Lancet Stillbirth Epidemiology investigator group. Stillbirths: rates, risk factors, and acceleration towards 2030. Lancet. 2016;387:587-603.
- 6. Fall CH, Sachdev HS, Osmond C, Restrepo-Mendez MC, Victora C, Martorell R, *et al.*; COHORTS investigators. Association between maternal age at childbirth and child and adult outcomes in the offspring: a prospective study in five low-income and middle-income countries (COHORTS collaboration). Lancet Glob Health. 2015; 3:e366-77.
- The State of the World's Children 2015: Executive Summary. http://www.unicef.org/publications/files/SOWC 2015_Summary_and_Tables.pdf. Accessed June 6, 2016.
- Markovitz BP, Cook R, Flick LH, Leet TL. Socioeconomic factors and adolescent pregnancy outcomes: Distinctions between neonatal and post-neonatal deaths? BMC Public Health. 2005;5:79.
- 9. Chen XK, Wen SW, Fleming N, Yang Q, Walker MC. Increased risks of neonatal and postneonatal mortality associated with teenage pregnancy had different explanations. J Clin Epidemiol. 2008;61:688-94.
- Kapoor RK, Srivastava AK, Misra PK, Sharma B, Thakur S, Srivastava KI, *et al*. Perinatal mortality in urban slums in Lucknow. Indian Pediatr. 1996;33:19-23.
- Raj A, Saggurti N, Winter M, Labonte A, Decker MR, Balaiah D, *et al.* The effect of maternal child marriage on morbidity and mortality of children under 5 in India: cross sectional study of a nationally representative sample. Br Med J. 2010;340:b4258.
- 12. Singh R, Tripathi V. Maternal factors contributing to under-five mortality at birth order 1 to 5 in India: a comprehensive multivariate study. Springerplus. 2013; 2:284.
- Restrepo-Méndez MC, Barros AJ, Santos IS, Menezes AM, Matijasevich A, Barros FC, *et al.* Childbearing during adolescence and offspring mortality: findings from three population-based cohorts in southern Brazil. BMC Public Health. 2011;11:781.
- 14. Richter LM, Victora CG, Hallal PC, Adair LS, Bhargava SK, Fall CH, *et al*; COHORTS Group. Cohort profile: the Consortium of Health-Orientated Research in Transition-

ing Societies. Int J Epidemiol. 2012;41:621-6.

- 15. Bhargava SK, Sachdev HS, Fall CH, Osmond C, Lakshmy R, Barker DJ, *et al.* Relation of serial changes in childhood body-mass index to impaired glucose tolerance in young adulthood. N Engl J Med. 2004;350:865-75.
- Bradburn MJ, Clark TG, Love SB, Altman DG. Survival analysis part II: multivariate data analysis — an introduction to concepts and methods. Br J Cancer. 2003;89:431-6.
- Vyas S, Kumaranayake L. Constructing socio-economic status indices: how to use principal components analysis. Health Policy Plan. 2006;21:459-68.
- Finlay JE, Ozaltin E, Canning D. The association of maternal age with infant mortality, child anthropometric failure, diarrhoea and anemia for first births: evidence from 55 low-and middle-income countries. BMJ Open. 2011;1:e000226.
- Haldre K, Rahu K, Karro H, Rahu M. Is a poor pregnancy outcome related to young maternal age? A study of teenagers in stonia during the period of major socioeconomic changes (from 1992 to 2002). Eur J Obstet Gynecol Reprod Biol. 2007;131:45-51.
- Chen XK, Wen SW, Fleming N, Demissie K, Rhoads GG, Walker M. Teenage pregnancy and adverse birth outcomes: a large population based retrospective cohort study. Int J Epidemiol. 2007;36:368-73.
- Gilbert W, Jandial D, Field N, Bigelow P, Danielsen B. Birth outcomes in teenage pregnancies. J Matern Fetal Neonatal Med. 2004;16:265-70.
- 22. Olausson PO, Cnattingius S, Haglund B. Teenage pregnancies and risk of late fetal death and infant mortality. Br J Obstet Gynaecol. 1999;106:116-21.
- Cowden AJ, Funkhouser E. Adolescent pregnancy, infant mortality, and source of payment for birth: Alabama residential live births, 1991-1994. J Adolesc Health. 2001;29:37-45.
- 24. Golding J, Greenwood R, McCaw-Binns A, Thomas P. Associations between social and environmental factors and perinatal mortality in Jamaica. Paediatr Perinat Epidemiol. 1994;8:17-39.
- 25. Were WM, Daelmans B, Bhutta Z, Duke T, Bahl R, Boschi-Pinto C, *et al.* Children's health priorities and interventions. Br Med J. 2015;351:h4300.
- 26. C-6. Ever Married and Currently Married Population by Age at Marriage, Duration of Marriage and Educational Level - 2011 (India and States/UTs/District level). Available from: http://www.censusindia.gov.in/2011 census/C-series/C-6.html(DDW-0000C-06.XLSX). Accessed June 6, 2016.
- NFHS 4 Factsheet. Available from: http://rchiips.org/ NFHS/factsheet_NFHS-4.shtm. Accessed March 30, 2016.

Variable		Stillbirth to under-five	five				Only perinatal	atal				Only Post-perinatal	-perinatı	al	
	Censored Alive (N= 5558)	d Alive 8)	Deaths (N=328	Deaths (N=328)	Ιd	Censored \neq (N= 5791)		De_{ℓ}	Deaths (N=95)	P2	Censored \neq (N= 5250)	Censored Alive (N= 5250)	Deaths N=233)	ths (33)	P3
	N	Mean (SD)%	N	Mean (SD)/%		Ν	Mean (SD)%	Ν	Mean (SD)%		N	Mean (SD)%	N	Mean (SD)%	
Child															
Birth order															
	5369	19.0	314	20.4	0.242	5591	18.9	92	27.2	0.216	5077	18.7	222	17.6	0.068
2		26.2		22.0			26.0		26.1			26.2		20.3	
3		21.8		20.4			21.7		19.6			21.7		20.7	
4+		33.0		37.3			33.4		27.2			33.4		41.4	
Gestation (wks)	4890	38.9 (2.7)	239	37.9 (3.4)	<0.001	5063	38.9 (2.7)	99	37.2 (4.0)	<0.001	4655	38.9 (2.7)	173	38.2 (3.2)	<0.001
Birth weight (g)	4644	2806 (427)	208	2398 (619)	<0.001	4814	2795 (437)	38	1968 (661)	<0.001	4338	2810 (426)	170	2494 (568)	<0.001
Birth length (cm)	4545	48.3 (2.2)	165	46.8 (3.1)	<0.001	4692	48.3 (2.2)	18	44.6 (4.2)	<0.001	4246	48.3 (2.2)	147	47.1 (2.9)	<0.001
Low Birth weight (<2500 g)	4644	24.8	208	54.8	<0.001	4814	25.7	38	71.1	<0.001	4338	24.3	170	51.2	<0.001
Predominantly breastfed at birth											3507	0.66	130	96.2	0.012
Predominantly breastfed at 3 months											3741	91.7	72	80.6	0.004
Mother															
Married	5555	100.0	326	100.0	1.000	5788	100.0	93	100.0	1.000	5247	100.0	233	100.0	1.000
Age (years)	5558	25.8 (5.2)	328	26.1 (6.1)	0.326	5791	25.9 (5.3)	95	25.5 (5.8)	0.473	5250	25.9 (5.2)	233	26.4 (6.2)	0.160
Education (formal years)	ars)														
Illiterate (0)	5548	35.2	327	52.0	<0.001	5781	36.1	94	38.3	0.043	5241	34.7	233	57.5	<0.001
Primary (1-5)		15.9		15.3			15.7		22.3			15.9		12.4	
Middle (6-10)		15.8		17.1			15.8		20.2			16.0		15.9	
Matric (up to 10)		20.6		12.2			20.3		14.9			20.9		11.2	
College (10 to 14)		12.4		3.4			12.1		4.3			12.5		3.0	
Place of delivery															
Home	5177	38.4	298	50.7	<0.001	5389	38.9	86	44.2	0.319	4870	38.4	212	53.3	<0.001
Healthcare service		61.6		49.3			61.1		55.8			61.6		46.7	
Family Demography	<i>iy</i>														
Household income 5553	5553	850.0 (2.1)	325	622.5 (1.9)	<0.001	5784	838 (2.1)	94	688 (1.9)	0.010	5245	850 (2.1)	231	598.0 (2.0)	< 0.001

SINHA, et al.

MATERNAL AGE AND OFFSPRING MORTALITY

INDIAN PEDIATRICS

VOLUME 53—OCTOBER 15, 2016

Variable		Stillbirth to under-five	r-five				Only perinatal	tal				Only Post-perinatal	-perinatu	11	
	Censored Alive (N= 5558)	d Alive 8)	Deaths (N=328	Deaths ($N=328$)	ΓI	Censored $(N=5791)$	Censored Alive (N= 5791)	Deaths (N=95)	ths 95)	P2	Censored $_{i}$ (N= 5250)	Censored Alive (N= 5250)	Deaths N=233)	ths (33)	P3
	N	Mean (SD)/%	×	Mean (SD)/%	'	N	Mean (SD)/%	N	Mean (SD)/%		~	Mean (SD)/%	N	Mean (SD)/%	
(per capita in Rs.)*	*_														
Type of housing															
Thatched hut	5543	1.6	320	4.1	<0.001	5771	1.8	92	0.0	0.114	5235	1.5	228	5.7	< 0.001
Masonry building		66.3		76.6			66.8		71.7			65.8		78.5	
Block of flats		27.7		18.4			27.2		28.3			28.1		14.5	
Bungalow		4.4		0.9			4.3		0.0			4.5		1.3	
Owned house	5542	40.8	319	38.2	0.380	5769	40.8	92	29.3	0.032	5234	41.8	227	41.9	1.000
Crowding	5182	3.6(1.8)	319	3.9 (2.0)	<0.001	5408	3.6(1.8)	93	3.5(1.9)	0.539	4874	3.6(1.8)	226	4.1 (2.1)	<0.001
Sanitation															
Open field	5188	22.2	321	37.4	< 0.001	5416	23.0	93	29.0	0.429	4880	21.6	228	40.8	< 0.001
Scavenger cleaned		37.4		38.0			37.5		32.3			37.9		40.4	
Pit		2.1		2.8			2.1		3.2			1.9		2.6	
Flush		38.3		21.8			37.4		35.5			38.5		16.2	
Water supply															
Unprotected	5547	17.9	320	25.6	<0.001	5775	18.3	92	16.3	0.649	5239	17.4	228	29.4	<0.001
Both (Unprotected & Protected)	_	9.0		15.9			9.4		12.0			8.8		17.5	
Protected		73.1		58.4			72.3		71.7			73.8		53.1	
Household wealth 5173 score (Standardized 'Z')	5173 ed 'Z')	0.03(1.00)	317	-0.44 (0.95) <0.001	<0.001	5398	0.002(1.0)	92	-0.09 (0.9) 0.403	0.403	4865	0.04 (0.99) 225) 225	-0.58 (0.92)	<0.001

SINHA, et al.

INDIAN PEDIATRICS

Web Table I continuted

Variables	Ν	Maternal age (per	decade)	Maternal age (per dec (quadratic term)	·
		Coefficient (95% CI)	P value	Coefficient (95% CI)	P value
Maternal education [#]	5875	2.96 (2.45; 3.48)	< 0.001	-0.59 (-0.68; -0.50)	< 0.001
Household income (₹)*	5878	0.36 (0.09; 0.62)	0.008	-0.12 (-0.16; -0.07)	< 0.001
Wealth ^{\$}	5490	1.58 (1.20; 1.95)	< 0.001	-0.27 (-0.34; -0.21)	< 0.001
Place of delivery (Healthcare services in comparison to home)	5475	7.95 (3.65; 17.31)	< 0.001	0.67 (0.58; 0.77)	< 0.001
Gestation (weeks)	5129	1.74 (0.67; 2.81)	0.001	-0.31 (-0.50; -0.12)	0.001
Birth weight (grams)	4852	736 (561; 912)	< 0.001	-116 (-147; -85)	< 0.001
Feeding at birth (only for post perinatal cases)	3637	0.77 (0.01; 49.78)	0.903	1.10 (0.54; 2.24)	0.795
Birth order	5683	4.51 (4.16; 4.86)	< 0.001	-0.61 (-0.68; -0.55)	< 0.001

WEB TABLE II	Association of Maternal Age as a Continuous Variable With the Available Confounders and Mediators
	(ADJUSTED FOR SEX)

[#]Maternal education categorized as 1- illiterate, 2- Primary, 3- Middle, 4- Matric and 5- College; * log transformed; [§]Household wealth was derived as 1st factor score generated from principal component analysis of type of housing, type of residence, sanitation, water supply and crowding (number of people/room).

SINHA, et al.

Variables	Model 1Hazard ratio (95% CI) (P value)	Model 2Hazard ratio (95% CI) (P value)	Model 3Hazard ratio (95% CI) (P value)
Mortality: All deaths			
Number of deaths/total sample (deaths + censored)	328/5886	316/5478	156/4154
Maternal age (per decade)	0.10 (0.03; 0.35) (<0.001)	0.12 (0.03; 0.46) (0.002)	0.13 (0.02; 0.80) (0.028)
Maternal age (per decade) (quadratic term)	1.52 (1.22; 1.90) (<0.001)	1.42 (1.12; 1.79) (0.003)	1.45 (1.07; 1.97) (0.018)
Sex (female in comparison to male)	1.21 (0.97; 1.50) (0.090)	1.19 (0.95; 1.48) (0.130)	1.00 (0.73; 1.37) (0.986)
Maternal education [#]		0.90 (0.80; 1.01) (0.077)	0.88 (0.75; 1.05) (0.150)
Household income (₹)*		0.70 (0.56; 0.86) (0.001)	0.69 (0.51; 0.94) (0.019)
Wealth ^{\$}		0.72 (0.63; 0.81) (<0.001)	0.74 (0.62; 0.88) (0.001)
Place of delivery (Healthcare services in con	nparison to home)		1.27 (0.90; 1.78) (0.174)
Gestation (weeks)			0.95 (0.90; 1.00) (0.062)
Birth weight (kg)			0.17 (0.11; 0.24) (<0.001)
Mortality: Perinatal deaths			
Number of deaths/total sample (deaths + censored)	95/5886	91/5478	29/4154
Maternal age (per decade)	0.10 (0.01; 1.23) (0.072)	0.10 (0.01; 1.39) (0.086)	0.21 (0.004; 10.59) (0.436)
Maternal age (per decade) (quadratic term)	1.46 (0.95; 2.26) (0.086)	1.43 (0.90; 2.72) (0.132)	1.35 (0.70; 2.58) (0.369)
Sex (female in comparison to male)	0.91 (0.61; 1.36) (0.643)	0.97 (0.65; 1.47) (0.901)	0.76 (0.36; 1.62) (0.476)
Maternal education [#]		0.94 (0.77; 1.15) (0.556)	0.99 (0.68; 1.44) (0.944)
Household income (₹)*		0.68 (0.46; 1.00) (0.050)	0.69 (0.34; 1.39) (0.296)
Wealth ^{\$}		1.07 (0.84; 1.34) (0.596)	1.20 (0.79; 1.83) (0.391)
Place of delivery (Healthcare services in con	nparison to home)		2.81 (1.03; 7.68) (0.045)
Gestation (weeks)			0.99 (0.86; 1.13) (0.826)
Birth weight (kg)			0.06 (0.02; 0.14) (<0.001)
Mortality: post-perinatal deaths			
Number of deaths/total sample (deaths + censored)	233/5483	225/5080	127/3894
Maternal age (per decade)	0.10 (0.02; 0.45) (0.003)	0.13 (0.03; 0.62) (0.011)	0.16 (0.02; 1.30) (0.086)
Maternal age (per decade) (quadratic term)	1.53 (1.18; 1.98) (0.001)	1.41 (1.08; 1.84) (0.013)	1.38 (0.97; 1.96) (0.071)
Sex (female in comparison to male)	1.36 (1.05; 1.76) (0.021)	1.28 (0.99; 1.67) (0.062)	1.07 (0.75; 1.53) (0.703)
Maternal education [#]		0.88 (0.77; 1.02) (0.080)	0.86 (0.71; 1.04) (0.115)
Household income (₹)*		0.70 (0.54; 0.90) (0.006)	0.72 (0.51; 1.01) (0.059)
Wealth ^{\$}		0.61 (0.53; 0.71) (<0.001)	0.66 (0.54; 0.80) (<0.001)
Place of delivery (Healthcare services in cor	nparison to home)		1.08 (0.75; 1.57) (0.685)
Gestation (weeks)			0.95 (0.89; 1.01) (0.110)
Birth weight (kg)			0.23 (0.15; 0.35) (<0.001)

WEB TABLE III ASSOCIATION BETWEEN	MATERNAL AGE AS A CONTINUOUS	VARIABLE AND MORTALITY
-----------------------------------	------------------------------	------------------------

*log transformed; [#]Maternal education categorized as 1- illiterate, 2- Primary, 3- Middle, 4- Matric and 5- College; ^{\$}Household wealth was derived as 1st factor score generated from principal component analysis of type of housing, type of residence, sanitation, water supply and crowding (number of people/room).

Model 1: adjusted for sex; Model 2: adjusted for sex, socio-economic confounders (maternal education, household income and wealth), Model 3: sex, socio-economic confounders and mediators (place of delivery, gestation and birth weight). The sample sizes in models varied because of completeness of data for all variables.